

| Substitute for form 1449/PTO <h2 style="text-align: center; margin: 10px 0;">INFORMATION DISCLOSURE STATEMENT BY APPLICANT</h2> <p style="text-align: center; margin: 10px 0;"><i>(Use as many sheets as necessary)</i></p> | | <table border="1" style="width: 100%; border-collapse: collapse;"> <tr> <th colspan="2" style="text-align: center; padding: 2px;">Complete if Known</th> </tr> <tr> <td style="width: 30%; padding: 2px;">Application Number</td> <td style="padding: 2px;">10/729,475-Conf. #3082</td> </tr> <tr> <td style="padding: 2px;">Filing Date</td> <td style="padding: 2px;">December 5, 2003</td> </tr> <tr> <td style="padding: 2px;">First Named Inventor</td> <td style="padding: 2px;">Steve PAKOLA</td> </tr> <tr> <td style="padding: 2px;">Art Unit</td> <td style="padding: 2px;">1651</td> </tr> <tr> <td style="padding: 2px;">Examiner Name</td> <td style="padding: 2px;">T. Kim</td> </tr> <tr> <td style="padding: 2px;">Attorney Docket Number</td> <td style="padding: 2px;">0113476.00122US1</td> </tr> </table> | | Complete if Known | | Application Number | 10/729,475-Conf. #3082 | Filing Date | December 5, 2003 | First Named Inventor | Steve PAKOLA | Art Unit | 1651 | Examiner Name | T. Kim | Attorney Docket Number | 0113476.00122US1 |
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| NON PATENT LITERATURE DOCUMENTS | | | |
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| | CA | Asami T, Terasaki H, Kachi S, Nakamura M, Yamamura K, Nabeshima T, and Miyake Y (2004) Ultrastructure of internal limiting membrane removed during plasmin-assisted vitrectomy from eyes with diabetic macular edema. <i>Ophthalmology</i> , 111, 231-237. | |
| | CB | Azzolini C, D'Angelo A, Maestranzi G, Codenotti M, Della Valle PPM, and Brancato R (2005) Intracavitary plasmin enzyme in diabetic macular edema. <i>Am J Ophthalmol</i> , 139, 949-950. | |
| | CC | Azzolini C, D'Angelo A, Maestranzi G, Codenotti M, Della Valle VP, Prati M, and Brancato R (2004) Intracavitary plasmin enzyme in diabetic macular edema. <i>Am J Ophthalmol</i> , 138, 560-566. | |
| | CD | Bhisitkul RB (2001) Anticipation for enzymatic vitreolysis. <i>Br J Ophthalmol</i> , 85, 1-2. | |
| | CE | Bishop PN (2000) Structural macromolecules and supramolecular organisation of the vitreous gel. <i>Prog Retin Eye Res</i> , 19, 323-344. | |
| | CF | Chen W, Huang X, Ma XW, Mo W, Wang WJ, and Song HY (2008) Enzymatic vitreolysis with recombinant microplasminogen and tissue plasminogen activator. <i>Eye</i> , 22, 300-307. | |
| | CG | Chow et al. "Successful Closure Of Traumatic Macular Holes" <i>The Journal of Retinal and Vitreous Diseases</i> , Vol. 19, No. 5, 1999 (405-409) | |
| | CH | Collen, D. "Revival of Plasmin as a Therapeutic Agent?" <i>Thromb Haemost</i> 2001; 86: 731-2 | |
| | CI | Foos, Robert Y. "Vitreoretinal junction: topographical variations" <i>Investigative Ophthalmology</i> , October 1972, Vol. 11, No. 10, (801-808) | |
| | CJ | Gandorfer A (2008) Experimental evaluation of microplasmin - an alternative to vital dyes. <i>Dev Ophthalmol</i> , 42, 153-159. | |

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| | CK | Gandorfer A and Kampik A (2005) Intrastretinal plasmin enzyme in diabetic macular edema. <i>Am J Ophthalmol</i> , 139, 949. | | |
| | CL | Goldberg GL, Strongin A, Collier IE, Genrich LT, and Marmer BL (1992) Interaction of 92-kDa type IV collagenase with the tissue inhibitor of metalloproteinases prevents dimerization, complex formation with interstitial collagenase, and activation of the proenzyme with stromelysin. <i>J Biol Chem</i> , 267, 4583-4591. | | |
| | CM | Hara A (1994) Surface structure of rabbit and human retina after enzymatic separation of inner limiting membrane. <i>Jpn J Ophthalmol</i> , 38, 375-381. | | |
| | CN | Hermel M and Schrage NF (2007) Efficacy of plasmin enzymes and chondroitinase ABC in creating posterior vitreous separation in the pig: a masked, placebo-controlled in vivo study. <i>Graefes Arch Clin Exp Ophthalmol</i> , 245, 399-406. | | |
| | CO | Jorge R, Oyamauchi EK, Cardillo JA, Gobbi A, Laicine EM, and Haddad A (2003) Intravitreal injection of dispase causes retinal hemorrhages in rabbit and human eyes. <i>Curr Eye Res</i> , 26, 107-112. | | |
| | CP | Joshi MM, Ciaccia S, Trese MT, and Capone A, Jr. (2006a) Posterior hyaloid contracture in pediatric vitreoretinopathies. <i>Retina</i> , 26, S38-S41. | | |
| | CQ | Joshi MM, Drenser K, Hartzler M, Dailey W, Capone A, Jr., and Trese MT (2006b) Intrachoroidal cavity fluid composition in congenital X-linked retinoschisis. <i>Retina</i> , 26, S57-S60. | | |
| | CR | Khono et al. "Immunofluorescent Studies of Fibronectin and Laminin in the Human Eye" <i>Investigate Ophthalmology & Visual Science</i> , vol. 28, March 1987, (506-514) | | |
| | CS | Kim NJ, Yu HG, Yu YS, and Chung H (2004) Long-term effect of plasmin on the vitreolysis in rabbit eyes. <i>Korean J Ophthalmol</i> , 18, 35-40. | | |
| | CT | Laryukhina GM and Ziangirova GG (1977) Experimental enzymatic vitreolysis. <i>Vestn Oftalmol</i> (6) 76-79. | | |

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| | CU | Li X, Shi X, and Fan J (2002) Posterior vitreous detachment with plasmin in the isolated human eye. Graefes Arch Clin Exp Ophthalmol, 240, 56-62. | |
| | CV | Marder et al. "Plasmin Induces Local Thrombolysis Without Causing Hemorrhage: A Comparison with Tissue Plasminogen Activator in the Rabbit" Thromb Haemost 2001; 86: 739-45 | |
| | CW | Moser TL, Enghild JJ, Pizzo SV, and Stack MS (1993) The extracellular matrix proteins laminin and fibronectin contain binding domains for human plasminogen and tissue plasminogen activator. J Biol Chem, 268, 18917-18923. | |
| | CX | Okajima K, Abe H, and Binder BR (1995) Endothelial cell injury induced by plasmin in vitro. J Lab Clin Med, 126, 377-384. | |
| | CY | Powell JR and Castellino FJ (1980) Activation of human neo-plasminogen-Val442 by urokinase and streptokinase and a kinetic characterization of neoplasmin-Val442. J Biol Chem, 255, 5329-5335. | |
| | CZ | Proksch JW, Driot JY, Vandeberg P, and Ward KW (2008) Nonclinical safety and pharmacokinetics of intravitreally administered human-derived plasmin in rabbits and minipigs. J Ocul Pharmacol Ther, 24, 320-332 | |
| | CA1 | Quiram PA, Leverenz VR, Baker RM, Dang L, Giblin FJ, and Trese MT (2007) Microplasmin-induced posterior vitreous detachment affects vitreous oxygen levels. Retina, 27, 1090-1096. | |
| | CB1 | Rizzo S, Pellegrini G, Benocci F, Belting C, Baicchi U, and Vispi M (2006) Autologous plasmin for pharmacologic vitreolysis prepared 1 hour before surgery. Retina, 26, 792-796. | |
| | CC1 | Sakuma et al. "Efficacy of autologous plasmin for idiopathic macular hole surgery" 2005 (Eur J Ophthalmol 15, 787-794) | |
| | CD1 | Sakuma T, Tanaka M, Inoue J, Mizota A, Souri M, and Ichinose A (2006) Use of autologous plasmin during vitrectomy for diabetic maculopathy. Eur J Ophthalmol, 16, 138-140. | |

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| | CE1 | Sebag J (1998) Pharmacologic vitreolysis. Retina, 18, 1-3. | | |
| | CF1 | Sebag J (2002) Is pharmacologic vitreolysis brewing Retina, 22, 1-3. | | |
| | CG1 | Sebag J (2005) Molecular biology of pharmacologic vitreolysis. Trans Am Ophthalmol Soc, 103, 473-494. | | |
| | CH1 | Sebag J and Balazs EA (1989) Morphology and ultrastructure of human vitreous fibers. Invest Ophthalmol Vis Sci, 30, 1867-1871. | | |
| | CI1 | Sebag J, Ansari RR, and Suh KI (2007) Pharmacologic vitreolysis with microplasmin increases vitreous diffusion coefficients. Graefes Arch Clin Exp Ophthalmol, 245, 576-580. | | |
| | CJ1 | Shapiro SD, Fliszar CJ, Broekelmann TJ, Mecham RP, Senior RM, and Welgus HG (1995) Activation of the 92-kDa gelatinase by stromelysin and 4-aminophenylmercuric acetate. Differential processing and stabilization of the carboxyl-terminal domain by tissue inhibitor of metalloproteinases (TIMP). J Biol Chem, 270, 6351-6356. | | |
| | CK1 | Stack MS, Moser TL, and Pizzo SV (1992) Binding of human plasminogen to basement-membrane (type IV) collagen. Biochem J, 284, 103-108. | | |
| | CL1 | Staubach F, Nuber V, and Janknecht P (2004) Enzyme-assisted vitrectomy in enucleated pig eyes: a comparison of hyaluronidase, chondroitinase, and plasmin. Curr Eye Res, 29, 261-268. | | |
| | CM1 | Takano A, Hirata A, Inomata Y, Kawaji T, Nakagawa K, Nagata S, and Tanihara H (2005) Intravitreal plasmin injection activates endogenous matrix metalloproteinase-2 in rabbit and human vitreous. Am J Ophthalmol, 140, 654-660. | | |
| | CN1 | Takano A, Hirata A, Ogasawara K, Sagara N, Inomata Y, Kawaji T, and Tanihara H (2006) Posterior vitreous detachment induced by nattokinase (subtilisin NAT): a novel enzyme for pharmacologic vitreolysis. Invest Ophthalmol Vis Sci, 47, 2075-2079. | | |

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| | CO1 | Trese MT (2000) Enzymatic vitreous surgery. Semin Ophthalmol, 15, 116-121. | | |
| | CP1 | Wang F, Wang Z, Sun X, Wang F, Xu X, and Zhang X (2004) Safety and efficacy of dispase and plasmin in pharmacologic vitreolysis. Invest Ophthalmol Vis Sci, 45, 3286-3290. | | |
| | CQ1 | Wang ZL, Zhang X, Xu X, Sun XD, and Wang F (2005) PVD following plasmin but not hyaluronidase: implications for combination pharmacologic vitreolysis therapy. Retina, 25, 38-43. | | |
| | CR1 | Wu HL, Shi GY, and Bender ML (1987) Preparation and purification of microplasmin. Proc Natl Acad Sci U S A, 84, 8292-8295. | | |
| | CS1 | Wu WC, Drenser KA, Capone A, Williams GA, and Trese MT (2007a) Plasmin enzyme-assisted vitreoretinal surgery in congenital X-linked retinoschisis: surgical techniques based on a new classification system. Retina, 27, 1079-1085. | | |
| | CT1 | Wu WC, Drenser KA, Trese MT, Williams GA, and Capone A (2007) Pediatric traumatic macular hole: results of autologous plasmin enzyme-assisted vitrectomy. Am J Ophthalmol, 144, 668-672. | | |
| | CU1 | Suenson, Elisabeth and Thorsen, Sixtus. "Secondary-site binding of Glu-plasmin, Lys-plasmin and miniplasmin to fibrin" Biochem. J. (1981) 197, 619-628 | | |
| | CV1 | Gonias, Steven and Figler, Nancy. " α 2-Macroglobulin is the primary inhibitor of miniplasmin <i>in vitro</i> and <i>in vivo</i> in the mouse" Biochem. J. (1988) 255, 725-730. | | |
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